

# BIODEGRADABLE SURGICAL IMPLANTS AND DEVICES

## Related Applications

5        This application is a continuation-in-part of U.S. Patent Application Serial Number 07/681,529, the entire disclosure of which are hereby incorporated by reference.

## Field of the Invention

10        The present invention relates to biodegradable surgical implants and/or devices.

## Background of the Invention

15        In surgery, it is known to employ at least partially biodegradable, elongated (typically tubular) surgical implants for supporting, connecting or separating elongated organs, tissues or parts thereof, such as canals, ducts, tubes, intestines, blood  
20        vessels, nerves, among others. In this context, a biodegradable, absorbable and/or resorbable, material refers to a material whose decomposition and/or dissolution products leave the system through metabolic ducts, kidneys, lungs, intestines and/or skin by secretion.

25        Examples of at least partially biodegradable implants include U.S. Patent No. 3,108,357 to Liebig which suggests a tubular device

comprises cylindrical, tubular, complementary parts.

However, known tubular, at least partially biodegradable surgical implants and devices involve several drawbacks and limitations. As for the implants including biostable parts, such as polymeric, and the like fibers, plastic or metallic coils or rings, or the like, such biostable parts or components remain in a patient's system even after a tissue or an organ has healed. Such components can later be harmful to a patient by causing infections, inflammatory reactions and like foreign matter reactions, and/or they might release particles, corrosion products, or the like which can wander in the system and/or cause harmful cellular level reactions.

Known tubular biodegradable implants manufactured by melt working technique or a like method are often massive and stiff and create, in resilient tissues, such as ducts, tubes, blood vessels, among others, an undesirable stiff, non-physiological bracing effect which can lead to harmful alterations in the properties of a tissue braced. In addition, the massive, tubular implants create a heavy local foreign matter loading the system at the installation site thereof and such loading can also contribute to harmful alterations in an operated tissue, such as canal, tube, duct, blood vessel, or the like.

On the other hand, the tubular structures constructed from

biodegradable fibers by braiding, knitting, weaving or some other similar technique do not possess the structural rigidity and/or resilience often required of a support implant to be fitted inside or outside a tubular tissue.

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### Summary of the Invention

It has been surprisingly discovered in the present invention that the deficiencies and drawbacks of known, at least partially biodegradable surgical implants and devices used for supporting, connecting or separating organs, tissues or parts thereof can be substantially eliminated with an implant, device or part thereof which is mainly characterized by comprising an at least partially biodegradable elongated member which is at least partially wound at least once around a center of rotation into a helical configuration and which is at least partially reinforced with biodegradable reinforcing elements.

An implant, device or part thereof (hereinbelow "device") according to the present invention can be conceived, having been formed in a manner that around a certain center point is wound some elongated member at a distance of a certain winding radius from the center point. If the winding center is stationary and the winding radius increases as the winding angle increases, the configuration of an obtained device is a spiral configuration, especially if the winding radius remains in the same plane. Provided that the

winding radius is constant and the winding center travels during the turning of an elongated member along a certain, for example, linear path, the device obtained has a circle-cylindrical screw-threaded configuration, or helix. On the other hand, if the winding radius changes while the winding center travels along a certain path, there will be produced a spiral configuration whose external surface is in conical shape. It is obvious that the implant, device or part thereof can include the above shapes and configurations as a combination and, for example, a combination of a spiral and a cylindrical screw-threaded configuration. The implant, device or part thereof can also be provided with members of other shapes in addition to a screw-threaded configuration, such as, for example, plates or sleeves.

The present invention relates also to a method for manufacturing an implant, device or part thereof ("device") including at least partially winding at least once an elongated blank formed by a biodegradable polymer matrix and biodegradable reinforcement elements.

The invention relates also to the use of an implant, a device and part thereof.

#### Brief Description of the Drawings

The present invention is described in more detail in the

following specification with reference made to the accompanying drawings. In the drawings,

Figure 1 represents a schematic perspective view of the formation of an array of lamellae turning into a fibrillated structure of a material micro-structure in an embodiment of a device of the invention;

Figure 2 represents an intra- and inter-fibrillary molecular structure;

Figure 3 represents a schematic view of the micro-structure of a fibrillated polymer;

Figure 4 represents a schematic view of the molecular structure of fibrillated devices according to embodiments of the present invention;

Figures 5 and 6 represent schematic perspective views of spiral-shaped embodiments of an embodiment of a device according to the present invention;

Figures 7a-9b represent schematic perspective views of conical embodiments of devices according to the present invention;

Figures 10a and 10b represent schematic perspective views of

a cylindrical embodiment of a device according to the present invention;

Figures 10c and 10d represent further embodiments of a  
5 cylindrical device according to the present invention;

Figures 11a-11l represent preferred cross sectional shapes and  
surface patterns for an elongated member, or blank, of embodiments  
of devices according to the present invention shown in Figures  
10 5-10;

Figures 12 and 13 represent a schematic perspective view of an  
embodiment of the invention;

Figure 14 represents a schematic view of an embodiment of a  
15 test arrangement according to the present invention described in  
example 1;

Figure 15 represents a surgical operation described in Example  
20 3;

Figure 16 represents a surgical operation described in Example  
4;

Figure 17 represents a schematic view of an embodiment of a  
25 device according to the present invention described in example 5;

Figure 18 represents another embodiment of a device according to the present invention;

Fig. 19 represents a cross-sectional view of an implant according to an embodiment of the present invention as it is being drawn;

Fig. 20 represents a cross-sectional view of the implant shown in Fig. 19 after being bent;

Fig. 21 represents a cross-sectional view of an implant according to an embodiment of the present invention at the time of implanting into a bile duct of a patient; and

Fig. 22 represents a cross-sectional view of the implant shown in Fig. 21 at some point after implantation, showing the expansion of the device.

#### Detailed Description of Preferred Embodiments

In the context of the present invention, the biodegradable reinforcing elements refer to the following:

- a) oriented or aligned structural units included in the micro-structure or molecular structure of a material such as oriented parts of molecules or parts thereof or

microfibrils, fibrils or the like oriented structural units formed thereby;

- 5           b) biodegradable organic filaments, fibers, membrane fibers or the like, or structures constructed thereof, such as bands, braids, yarns, fabrics, non-woven structures or the like; or
- c) biodegradable inorganic (ceramic) filaments, fibers, membrane fibers or the like, or structures constructed thereof.

10           A particularly preferred embodiment of the present invention is such an implant or device which is structurally self-reinforced. A self-reinforced biodegradable structure is defined in U.S. Patent No. 4,743,257 to Törmälä et al. In a self-reinforced structure, a  
15           biodegradable polymer matrix is reinforced with biodegradable reinforcement elements or units having the same proportional elemental composition as the matrix. The reinforcement elements are typically oriented molecules or parts thereof or the like obtained by orientation of fibrils, microfibril, fibers, filaments  
20           or the like structures constructed thereof.

          Reinforcement elements inside the microstructure of a self-reinforced polymer material are produced, for example, by orienting the molecular structure of a material either in melt  
25           state or in solid state. In such conditions, the structure-reinforcing orientation remains at least partially, permanently in



material either as a result of the rapid cooling and/or solid state of the melt and/or as a result of the prevention of molecular movements (relaxation) of the melt. The self-reinforcement based on draw orientation is described in the invention PCT/FI87/00177, 5 Törmälä et al., as follows.

A partially crystalline, non-oriented piece of polymer typically consists of crystal units, that is, spherulites and amorphous areas thereinside and/or therebetween.

10 The orientation and fibrillation of a polymer system possessing a spherulitic crystalline structure is a process that has been extensively studied in connection with the production of thermoplastic fibers. For example, U.S. Patent 3,161,709 suggests 15 a three-step drawing process for transforming a melt-worked polypropene filament into a fiber having a high tensile strength.

20 The mechanism of orientation and fibrillation is basically as follows (C.L. Choy et al. Polym. Eng. 30 Sci. 23, 1983, p. 910). As a partially crystalline polymer is being drawn, the molecular chains of crystal lamellae quickly begin to parallel, or orient, themselves in the drawing direction. In other words, the reinforcement elements lie substantially parallel to the longitudinal axis of the rod. The drawing orients the 25 reinforcement elements to the longitudinal direction of the rod. The first phase of the drawing of an implant according to an

embodiment of the present invention is illustrated in Figure 19. As can be seen in Figure 19, the fibers of the matrix are oriented in the longitudinal direction of the device, the same direction as the drawing, which is indicated by the arrows.

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Simultaneously, the spherulites extend in length and finally break. Crystal blocks detach from lamellae and join together as queues by means of tight tie-molecules which are formed through the partial release of polymer chains from crystal lamellae. The alternating amorphous and crystalline zones, together with tight tie-molecules, form long, thin approximately 100 Å wide microfibril which are paralleled in the drawing direction. Since the intrafibrillar tie-molecules form in the phase boundaries between crystal blocks, they will be mainly located on the external surface of microfibrils. Those tie-molecules, which link various lamellae in an isotropic material prior to the drawing, serve in a fibrillated material to link various microfibrils together, that is, become interfibrillar tie-molecules which are located in boundary layers between adjacent microfibrils.

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The rod is then bent by winding to a helical configuration. The bending actually creates a "pre-stress" condition to the wound rod which then tends to open the helix beyond the diameter it was wound. Figure 20 illustrates the bending and the forces on the implant. The arrows to the right and left of the implant indicate the direction the implant tends to move in response to the bending.

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In other words, the arrows represent an opening force.

Figure 1 illustrates, schematically, the transformation of an array of lamellae into a fibrillar structure (a fibril consisting of a bunch of microfibrils) due to the action of water. Figure 2 shows some of the molecular structure inside and between microfibrils. Figure 3 illustrates, schematically, some of the structure of a fibrillated polymer. The Figure shows several fibrils one being dyed grey for the sake of clarity, which comprise a plurality of microfibrils having a length of several microns.

Orientation is initiated right at the start of drawing and also a fibrillated structure is formed at rather low drawing ratios  $\lambda$ , wherein  $\lambda = (\text{length of a piece after drawing})/(\text{length of piece prior to drawing})$ . For example, HD-polyethene is clearly fibrillated at  $\lambda$  value 8 and polyacetal (POM) at  $\lambda$  value 3.

As the drawing of a fibrillated structure is continued further (this stage of the process is after referred to as ultra-orientation), the structure is further deformed with microfibrils sliding relative to each other to further increase the proportional volume of straightened interfibrillar tie-molecules. If the drawing is effected at a sufficiently high temperature, the oriented tie-molecules crystallize and build axial crystalline bridges which link together crystalline blocks.

The excellent strength and modulus of elasticity properties of a fibrillated structure are based on the vigorous orientation of polymer molecules and polymer segments in the direction of drawing (in the direction of the longitudinal axis of microfibrils) characteristic of the structure.

The fibrillation of macroscopic polymeric blanks, such as rods or tubes, is prior known in the cases of biostable polyacetal and polyethylene (See K. Nakagawa and T. Konaka, Polymer 27, 1986, p. 1553 and references included therein). What has not been prior known, however, is the orientation and fibrillation of at least partially helical and/or spiral or similarly shaped members or pieces manufactured from biodegradable polymers.

The at least partial orientation and/or fibrillation of a biodegradable helical and/or spiral or similar piece can be effected, for example, by rapidly chilling a flowing state polymer melt, for example, in an injection mold, into a solid state in a manner that the orientation of molecules existing in the flowing melt in flowing direction is not allowed to discharge through molecular movements either entirely or partially into a state of random orientation.

A more vigorous orientation and fibrillation and, thus, also improved mechanical qualities are generally provided for a polymer piece by mechanically working the material (orientation), and

generally drawing or hydrostatic extrusion or die-drawing in such a physical condition (usually in solid state). Under these conditions, it is possible for the material to undergo dramatic structural deformations in its crystalline structures and amorphous areas occurring at the molecular level for creating orientation and fibrillation. As a result of fibrillation, for example, a restorable polymer material produced by injection molding or extrusion and initially possessing mainly a spherulitic crystalline structure, transforms into a fibrillated structure which is vigorously oriented in the direction of drawing and comprises, for example, elongated crystalline microfibrils as well as tie-molecules linking them as well as oriented amorphous areas. In a partially fibrillated structure, the amorphous areas between microfibrils make up a more substantial portion of the material than in an ultra-oriented material which, in most preferred case, only includes amorphousness as crystal defects. As a result of orientation, fibrillation and ultra-orientation, the values of strength and modulus of elasticity of a material are multiplied compared to a non-fibrillated structure.

Orientation and the resulting fibrillation can be used for treating biodegradable polymers, copolymers and polymer compositions so as to form self-reinforced composites in which nearly the entire material stock is oriented in a desired fashion and the portion of amorphous matrix is small. The stock orientation and small amount of matrix result in such materials

having extremely high quality strength properties in orientation direction with a bending strength, for example, up to 400-1500 MPa and modulus of elasticity 20-50 GPa. Thus, the orientation and fibrillation can be used to provide helixes or spirals or the like devices with multiple strength values compared to those of normal melt-processed biodegradable materials, which are typically in the order of 30-80 MPa.

As in the fibrillated structure of polymer fibers, in the structure of fibrillated devices, there can be found, for example, the following structural units, which are schematically shown in Figure 4: crystalline blocks, the stock therebetween comprising an amorphous material, for example, loose polymer chains, chain ends and molecular folds; tie-molecules which link the crystalline blocks together, the number and tightness of these increases as drawing ratio  $\lambda$  increases; as well as possible crystalline bridges between crystalline blocks. Bridges can form during the drawing as tie-molecules orient and group themselves as bridges (C.L. Choy et al., J. Polym. Sci., Polym. Phys. Ed., 19, 1981, p. 335).

The oriented fibrillated structure shown in Figures 1-4 is already developed by using so-called "natural" drawing ratios 3-8. As drawing is then continued as ultra-orientation, the portion of crystalline bridges can increase to be quite considerable whereby, in the extreme case, the bridges and crystal blocks provide a continuous crystalline structure. However, the effects of tie-

molecules and bridges are often similar and, thus, the exact distinction thereof from each other is not always possible.

Orientation and fibrillation can be experimentally characterized by the application of several different methods. Orientation function ( $f_c$ ), which can be determined by X-ray diffraction measurements, characterizes orientation of the molecular chains of a crystalline phase. Generally,  $f_c$  already reaches a maximum value of 1 by natural drawing ratios ( $\lambda < 6$ ). For polymer materials having a spherulitic structure  $f_c \ll 1$ .

Double-refraction ( $\Delta$ ) measured with a polarization microscope is also a quantity which represents the orientation of molecular chains. It generally increases at natural drawing ratios ( $\lambda < 6$ ) vigorously and, thereafter, in ultra-orientation more slowly, which indicates that the molecular chains of a crystalline phase orient vigorously in the drawing direction at natural drawing ratios and orientation of the molecules of an amorphous phase continues further at higher drawing ratios (C.L. Choy et al., Polym Eng. Sci., 23, 1983, p. 910).

The formation of a fibrillous structure can also be demonstrated visually by studying the fibrillated material by means of optical and/or electrical microscopy (see e.g. T. Knoda et al., Polymer, 26, 1985, p. 462). Even the individual fibrils consisting of microfibrils can be clearly distinguished in scanning electron

microscope images of a fibrillated structure.

5 An oriented and/or fibrillated and/or ultra-oriented piece (blank) is then rotated at least once around a center of rotation into helical configuration to form "a device" of the invention by shaping the blank at least partially by means of an external force or pressure and/or external heat and/or by means of heat inducible in the piece, for example, by radiowave radiation. In practice, the rotation of winding of the device is effected in a manner that an elongated blank is wound around a suitable, if necessary heated mold, for example, a mold of cylindrical shape). Such a mold is typically round in cross-section so as to produce helical shapes having a circular cross-section. The cross-sectional shape of a mold can also be, for example, elliptical, oval, or angular to produce helical shapes having various cross-sections. Orientation, fibrillation or ultra-orientation can also be effected in a continuous action and/or simultaneously with winding in a manner that an elongated blank is being drawn and the drawn section is simultaneously wound around a cylindrical mold.

20 At least partially oriented and/or fibrillated and, particularly, ultra-oriented biodegradable devices are an example of an oriented, self-reinforced biodegradable (U.S. Patent 4,743,257, Törmälä et al.) composite material, wherein the oriented reinforcement elements, such as fibrils, microfibrils, crystal blocks, tie-molecules or crystallized bridges, among others, are



formed and/or grouped during a mechanical working and a phase binding. The oriented reinforcement elements comprise, for example, of the following structural elements: amorphous phase, interfaces between crystal blocks as well as interfaces between bridges and microfibrils, a typical feature of which is also a vigorous orientation in a drawing direction.

Another method for using biodegradable reinforcement elements in devices of the invention is the reinforcement thereof with fibers manufactured from polymer, copolymer or a polymer composition, with film fibers, filaments or structures constructed thereof, such as braids, threads, ribbons, non-woven structures, fabrics, knittings or the like, by combining readymade fibers with a suitable polymer matrix. Such fibers can be manufactured, for example, from biodegradable polymers set forth in Table 1. The fibers can also be biodegradable ceramic fibers, such as calcium phosphate fibers (see e.g. S. Vainionpää et al., Progr. Polym. Sci., in printing).

Various plastic technological methods can be applied to manufacture devices of the present invention reinforced with biodegradable organic and/or inorganic fibers or with structures constructed thereof. The manufacturing may be carried out by binding the reinforcement structures at least partially to each other with biodegradable polymer, copolymer or a polymer composition (matrix) in such conditions which serve to produce a

sufficiently equal quality composite from the matrix and reinforcement elements. The matrix is usually in solution or melt state. Methods for combining reinforcement fibers or the like and a matrix as well as for processing them into semi-finished products and/or devices include, among others, injection molding, extrusion, pultrusion, winding, and compression molding.

An at least partially spirally shaped, at least partially biodegradable device according to the present invention can be used in a versatile manner for supporting, expanding, joining or separating organs, tissues or parts thereof. A device according to the present invention offers a plurality of benefits over the prior art implants and devices. When using a device according to the invention, the amount of foreign matter remains smaller than with traditional implant tubes. Devices according to the present invention are more flexible and resilient than the rigid prior art tubes and, on the other hand, devices according to the present invention are stronger under compression and retain their shape better than fiber-constructed tubular devices, whereby devices according to the present invention are capable of being used for retaining open or even expanding the medullary cavity of tubular tissues.

Devices of the present invention can be manufactured from biodegradable polymers, copolymers and polymer compositions. Table 1 shows a number of prior known biodegradable polymers, which or

29. Poly- $\beta$ -maleic acid (PMLA)
30. Poly- $\beta$ -alkanoic acids
31. Polyethylene oxide (PEO)
32. Chitin polymers

Reference: S. Vainionpää, P. Rokkanen and P. Törmälä, Progr. Polym. Sci., in printing

It is obvious that biodegradable polymers other than those set forth in Table 1 can also be used as raw materials for implants, devices or parts thereof. For example, the biodegradable, absorbable polymers described in the following publications can be used for the above purposes: U.S. Patent No. 4,700,704 to Jamiolkows and Shalaby; U.S. Patent No. 4,655,497 to Bezwada, Shalaby and Newman; U.S. Patent No. 4,649,921 to Koelmel, Jamiolkows and Bezewada; U.S. Patent No. 4,559,945 to Koelmel and Shalaby; U.S. Patent No. 4,532,928 to Rezada, Shalaby and Jamiolkows; U.S. Patent No. 4,605,730 to Shalaby and Jamiolkows; U.S. Patent No. 4,441,496 to Shalaby and Koelmel; U.S. Patent No. 4,435,590 to Shalaby and Jamiolkows; and U.S. Patent No. 4,559,945 to Koelmel and Shalaby.

It is also natural that devices of the present invention may contain various additives and adjuvants for facilitating the processability of the material such as, for example, stabilizers, antioxidants or plasticizers; for modifying the properties thereof such as, for example, plasticizers or powdered ceramic materials or biostable fibers such as, for example, carbon fibers; or for facilitating the manipulation thereof such as, for example, colorants.

According to one preferred embodiment, devices of the present invention may contain some bioactive agent or agents, such as antibiotics, chemotherapeutic agents, wound-healing agents, growth hormone, contraceptive agent, anticoagulant, such as heparin. Such bioactive devices are particularly preferred in clinical applications since, in addition to mechanical effect, they have biochemical, medical and the like effects in various tissues.

Devices of the present invention can also be advantageously combined with other types of biodegradable implants and devices. For example, by inserting a helical device as shown in Figures 7-10 into a tube woven or knitted from biodegradable and/or biostable thread there is obtained a firm and resilient tube which has a variety of applications in surgery for replacing or supporting tissues and/or for keeping open the cavities within or between tissues.

A device according to the present invention can also be fitted with long biodegradable rods which extend parallel to the longitudinal axis of, for example, a helically-shaped device. Thus, if necessary, the device can be braced to form a tubular structure.

A device according to the invention can also be fitted with various other accessories, such as flat, perforated plates at the ends of a device for securing the ends of a device firmly to the

surrounding tissues by means of surgical stitches.

Devices according to the present invention can have various geometrical configurations. For example, Figure 5 illustrates a flat or planar (in imaginary plane 1) spiral 2 that can be used as a resilient separating material between tissues. The helixes of spiral 2 can also be connected with each other by means of biodegradable radial wires, rods 3 or the like as shown in Figure 6. The spiral has a high strength in the direction of plane 1 but is resilient in the direction perpendicular to that plane.

A device according to the present invention can also vary in its dimensions in various sections thereof. For example, Figures 7a-10a and 7b-10b schematically illustrate a few such devices. These devices may be used for providing external and/or internal support for organs or their parts of various shapes such as liver, spleen, kidneys, intestines, among others.

A device shown in Figure 7a is wound into a conical body. The conical body can have a side face outline which is either straight or arched or a combination thereof according to the intended application. Devices shown in Figures 8a and 9a include two conical bodies joined to each other either at the base of conical bodies as shown in Figure 8a, or at the apex thereof, as shown in Figure 9a. Figure 10 illustrates a device having its outer face wound into a cylindrical configuration.

Figures 7b, 8b, 9b and 10b illustrate device configurations matching those of Figures 7a, 8a, 9a and 10a and fitted rods 3 connecting the turns of helical body.

5 Furthermore, Figure 10c shows an embodiment in which the device is comprised of two nested device elements VO1 and VO2 wound into a helical configuration preferably in opposite directions. Each has a cylindrical helical configuration. Figure 10d shows an embodiment of a device, wherein a number of device elements wound  
10 into a helical configuration have been twined together. The device elements are adapted to run alternately over and under each other to form a tubular structure.

15 Figures 11a-11l illustrate some types of cross-sections for a blank. A blank for manufacturing devices of the present invention can have a cross-section which is, for example, circular (11a), elliptical (11b, 11c), flat (11d, 11e), angular (11f, 11g, 11h) or asteroid (11i). By varying the cross-section of a blank it is possible to effect, for example, on the mechanical properties of a  
20 device, the growth of tissues on the surface of a blank and the growth of tissues through the device. The thickness of a blank can also vary in different sections of a blank or it can be provided with holes R (11j) or similar structures, such as recesses L (11k) or slots (11l) for facilitating the fastening or securing thereof  
25 to tissues.

According to one preferred embodiment, a device of the present invention is manufactured by winding or rolling a flat blank having a cross-section shown in Figure 12 into a tube as shown in Figure 13. Since the longitudinal edges of a blank as shown in Figures 12a and 13a are provided with folded or other gripping means T which engage each other, during the winding, there will be formed a flexible tube that can be used in the treatment of, for example, a windpipe or the like flexible tissue channels as a temporary prosthesis.

According to one preferred embodiment, devices of the present invention can be used to join together tissues, organs or parts thereof, such as muscular tissue or the other soft tissues. Such an embodiment is illustrated in Figure 18. Figure 18a shows a cross-section of a tissue K1 and a tissue K2 which should be joined with each other. The joining can be effected by using a sharp-pointed spiral S which is driven the same way as a corkscrew through the tissues (Figure 18b). By locking the top portion of a spiral in position after the turning, for example, by stitching the spiral firmly to surrounding tissues by means of surgical stitches dissolving through the holes made in the spiral blank, the spiral serves to secure tissues K1 and K2 to each other preventing the separation or sliding thereof relative to each other.

Unexpectedly, embodiments of the present invention will expand upon implantation into tissue. The expansion may, therefore,

provide the surprising advantage of helping to lock the implant in place. This expansion is the result, at least in part, of the composition and manufacturing technique of the invention and of the reaction of the invention to the body temperatures to which the invention may be subjected to after implantation. The expansion of the implant may be at least about 15% when it is implanted in conditions of living tissue.

The present invention and its applicability is described in more detail by means of the following examples.

#### EXAMPLE 1.

Some polymers set forth in Table 1 were used to prepare helical devices of the present invention. An example, such as that shown in Figure 10, includes blank thickness 1 mm, outer diameter of helix 6 mm, inner diameter 4 mm, pitch angle 15 degrees and length of device 15-20 mm. The polymeric melt is subjected to injection molding to produce blanks having a diameter ( $\phi$ ) of 1.5 - 2.0 mm by drawing (orientation and self-reinforcement) then at a temperature of  $T_M > T > T_g$ , wherein  $T_g$  is polymer glazing temperature and  $T_m$  is polymer (possibly) melting temperature, to the  $\phi$  reading of 1 mm and by winding them in hot state around a metal pipe of diameter 4 mm. The device is then cooled and the finished device is removed from the surface of the metal pipe.



Reference materials were made by using similar polymers to prepare tubular pieces including a tube length of 10 mm, outer diameter of 6 mm and inner diameter of 4 mm by injection molding polymer melt into cooled tubular mold. The compression strength of the devices and that of the corresponding tubes were compared to each other by squeezing a device (Figure 14b) or a tube (Figure 14a) placed between two steel plates with an external force in the direction orthogonal to its longitudinal axis. The bending of a device in lateral direction was prevented by prepressing the device into a compact bundle between two vertical plates (Figure 14b).

The compression load strengths of a tube (Figure 14a) a device (Figure 14b) made of the same polymer and having equal weights were compared to each other. This was followed by the determination of the relative compression load strength (SP) of the device using the formula: (force required to fracture the device)/(force required to fracture the tube). Devices and tubes were manufactured by using the following biodegradable polymers, copolymers and polymer compositions: polyglycolide (Mw 60,000), glycolide/lactide copolymer (Mw 40,000), glycolide/trimethylenecarbonate copolymer (Mw 60,000), PLLA (Mw 260,000), PDLLA (Mw 100,000), lactide/ $\delta$ -valerolactone copolymer (Mw 60,000), lactide/ $\epsilon$ -capro-lactone copolymer (Mw 60,000), PHBA (Mw 700,000), PHPA (Mw 50,000) and PDS (Mw 40,000). Resulting values for SP were ranging between 1.8-12.

#### EXAMPLE 2.

times for preparing the device. A corresponding device was made by using just polyurethane.

Devices corresponding to such reinforced devices were also manufactured from mere thermoplastic matrix polymers by the application of melt working technique.

Table 2 illustrates the matrix polymers and fibrous reinforcements for the devices prepared.

Table 2. Structural components for fiber-reinforced biodegradable devices.

<u>Matrix polymer</u>	<u>Fiber reinforcement</u>
PDS	PGA
- "-	PGA/TMC
- "-	PGA/PLLA
- "-	PLLA
- "-	PHBA
- "-	PHBA/HVA
- "-	Chitin fiber
- "-	PDS
PDLLA	PGA
- "-	PGA/TMC
- "-	PGA/PLLA
- "-	PLLA
- "-	PHBA
- "-	PHBA/HVA
- "-	PDS
- "-	PDLLA
PLLA	PGA
- "-	PGA/TMC
- "-	PLLA
PVA	PGA
- "-	PGA/TMC
- "-	PGA/PLLA
- "-	PLLA
- "-	PHBA
- "-	PHBA/HVA

	- "-	PDS
	- "-	Chitin fibres
5	PGA/TMC	PGA
	- "-	PGA/TMC
	PHBA	PGA
	- "-	PGA/TMC
10	- "-	PHBA
	Poly-ε-caprolactone	PGA
	- "-	PGA/TMC
	- "-	PHBA
15	Methylmetacrylate- N-vinylpyrrolidone	PGA
	Polyurethane	PGA
20		Collagen (catgut)
	PEO	PGA
	- "-	PGA/TMC
	- "-	PGA/PLA
25	- "-	PLLA
	n-Butylcyano- acrylate	Collagen (catgut)
		PGA

The devices were secured by their ends to a tension apparatus and were drawn until broken in the direction of longitudinal axis of the device which corresponds to the winding axis of the blank. This was followed by the determination of the relative tensile load-bearing strength (SV) of a reinforced device using the formula: (force required to fracture a reinforced device)/(force required to fracture a corresponding non-reinforced device). The SV values were ranging between 1.5-8.

### EXAMPLE 3.

Preparation of a self-reinforced polyactide device as shown in

Figure 10 (hereinbelow "helix" (KR)) was formed using a raw material of poly-L-lactide/poly-DL-lactide copolymer (PLLA/PDLLA molar ratio 80/20, Mw 60,000). The helix (KR) was manufactured from a thick, extrusion-made PLLA/PDLLA rod, which was drawn to a drawing ratio of  $\lambda=7$  at a temperature of 90 degrees for self-reinforcing the material. A thus prepared self-reinforced rod having a thickness of 1 mm was then wound to form "a helix" as described in Example 1. The helix was cut into lengths of 12 mm for the following examination.

Under general anaesthesia, the gastric cavity of a dog was opened, the intestines were set aside and the bile duct (ST) was exposed by preparation, (see Figure 15). A roughly 6 mm long incision (AK1) was made in the duct. As shown in Figure 15a, a distance of 5 mm of this incision was provided with non-resorbable stitches (KO), which pucker up the duct and narrow it permanently together with a cicatricial tissue formed on incision (AK1). This was followed by closing the gastric cavity, stitching the skin and, after waking up from the anaesthesia, the dog was allowed to move freely in its cage.

After one month, the dog was re-anaesthetized, the gastric cavity was incised and the blocked bile duct was prepared to re-expose it. The duct was opened with a longitudinal incision (AK2) at the region of cicatricial pucker and a helix having an inner diameter of 2 mm and an outer diameter of 3 mm was inserted.

The helix was inserted such that both of its ends were located in healthy bile duct and its central portion within the incised pucker region. The bile duct was closed with a stitch (O), whereby its walls extended around the spiral. The situation is schematically  
5 illustrated in Figures 15c, 21 and 22.

After the operation, the bile duct was normal in volume. In other words, the implant has expanded. Such an expansion of, in this example, the bile duct is a manifestation of the unexpected  
10 result that based, at least in part upon the manufacturing technique of the present invention and the reaction of the present invention to body temperatures the present invention is subjected to after an operation, the biodegradable implant of the present  
15 invention may expand upon implantation in tissue. This expansion will lock the implant in place. Figures 21 and 22 illustrate this aspect of the invention. Figure 21 represents the implant at the time of implantation and Figure 22 represents the implant after expansion due to the body conditions has caused the implant to expand. If the implant did not expand, the duct would remain as  
20 shown in Figure 21, with the narrower portion defined by the implant. The gastric cavity and skin were closed the same way as in the first operation. The dog was put away after 14 months by which time the helix had nearly disappeared and the bile duct had a normal extent and volume and the pucker was no longer  
25 macroscopically observable.

#### EXAMPLE 4

Under general anaesthesia the right hind femoral vein (RL in Figure 16a) of a dog was cut. The base portion of the more distal vein was threaded into the interior of a biodegradable, reinforced device ("helix") having an inner diameter of 8 mm, an outer diameter of 9 mm and a length of 2 cm. The device, being like the one shown in Figure 10 (reinforcing fibers: Ca/P-fibers; matrix polymer PLLA, Mw 100,000; fiber/polymer weight ratio = 30/70 (w/w) and vein (LO), was stitched with end-to-end technique by using a resorbable 6-0 yarn to make a tight seam with no bleeding. After the operation, due to the flabbiness of the walls, the vein tended to collapse within the region of the stitched seam. This leads to a poorer circulation in the vein resulting easily in the development of a coagulation or clot formed by blood particles within the region of the seam and, thus, the veins will be blocked. The situation is illustrated in the schematic view of Figure 16a from the side of stitched seam, and in Figure 16b from above a stitched seam. Therefore, a biodegradable helix (KR) was pulled over the seam portion with the stitched seam remaining at the half-way point of helix (KR). The wall of a vein was attached at the stitched seam over its entire circumference to the helix by means of non-restorable support stitches (TO) (Figure 16c). This way, the vein was tensioned to its normal extent with the help of a support provided by the device. After the seam had healed, especially after the inner surface of a blood vessel or endothelium

had healed, there is no longer a risk of developing a clot and helix can resorb away with no harm done. After 6 months, the dog was put away and the femoral vein had healed with a pucker or clot.

5      EXAMPLE 5.

10      The test animals used in this example were male rabbits weighing 3 kg. The animals were anesthetized for the operation with im. ketamin and iv. pentobarbital preparations. A polylactide blank ( $\phi$  1 mm) was used to prepare a helix having an outer diameter of 8 mm and a length of 15 mm and an extension formed by a thin tubular neck section, 10 mm, followed by two helical coils (Figure 17).

15      The anesthetized test animals were subjected to a surgical incision of the urinary bladder through abdominal covers. Through the opened bladder, the prosthesis was threaded into position with the narrow neck section remaining within the region of closure muscle and the helical coil ends on the side of the urinary bladder.  
20

25      The prothesis was fitted in 15 test animals which were under observation for 3 months. The study verified that an implant of the invention can be used for preventing a lower urethra obstruction caused by the enlargement of forebland.

EXAMPLE 6.

The test animals in this example were male rabbits weighing approximately 3 kg. The animals were anesthetized for the operation with im. ketamin and iv. pentobarbital preparations.

The implants employed were PLLA helixes as described in Example 1 (cross-section of blank circular, thickness of blank 1 mm, outer diameter of helix 6 mm and length 15 mm).

On the anesthetized test animals was performed scission of the blind urethra to the extent sufficient for a prothesis. The prothesis was placed on the distal side of closure muscle. In connection with the operation an antibiotic was administered as a single dose: ampicillin 100 mg/kg.

The prothesis was fitted in 15 animals which were put away with iv overdose of anesthetic 2 weeks, 3 months, 6 months, 1 year and 2 years after the implantation. The urethra was dissected and tissue samples were taken for histological and electron microscopic analysis.

Histological studies indicated that PLLA had caused only slight foreign matter reaction in tissues. Two years after the implantation the helix had nearly completely biodegraded and the urethra was almost normal in its dimensions(i.e. the implant has



expanded). This state of the bile duct would not be possible if the bile duct had not expanded at the helix. If the helix had not expanded, there would be a constriction in the duct corresponding to the thickness of the helix wall, as shown in Figure 21.

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#### EXAMPLE 7.

Cloggings in the ureters leading from kidney to bladder will become more common as a result of the increased observation surgery of upper urethras. The ureter has a good regeneration ability when subjected longitudinal incision but its healing requires an internal support. Transverse incision or short deficiency always leads to the development of a clogging.

The purpose of this example was to examine the applicability of a helix made of a biodegradable material both the healing of a longitudinal dissection of the urethra and to the healing of a transverse deficiency.

The test animals were female rabbits weighing approximately 3 kg. The animals were anesthetized. Incision of the abdominal cavity was performed on the flank without opening, however, the actual abdominal cavity.

a) the urethra having a diameter of ca. 4 mm was dissected lengthwise over a distance of ca. 2 cm followed by threading

a self-reinforced PGA-helix (blank thickness 1 mm) inside the urethra, the helix having an outer diameter of 4 mm and a length of 20 mm. The region of dissection was covered with fat.

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- b) a length of ca. 1 cm was cut off the urethra, the remaining ends were dissected over a distance of 0.5 cm and the above-described prothesis was threaded in, so that the remaining defect zone of the tissue was 1 cm. The defect zone was covered with surrounding fat. After 1 month, 3 months and 1 year from the operation a tracer imaging of the kidneys was performed for observing the healing of the urethra. The operated urethras had healed to almost normal condition over the period of 1 year (on the basis of tracer imaging).

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In this disclosure, there is shown and described only the preferred embodiments of the invention, but, as aforementioned it is to be understood that the invention is capable of use in various other combinations and environments and is capable of changes or modifications within the scope of the inventive concept as expressed herein.